Due Date: June 14, 2003 (Saturday)

Art Unit: 1642

Examiner: Harris, A.M.

Docket: 0609.4560002

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Exhibi+A 09/720,086

Application No.: 09/720,086

Applicant:

102(e): July 23, 2001

Li et al.

Atty: JAG/KRM/DJN

For: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

When receipt stamp is placed hereon, the USPTO acknowledges receipt of the following documents:

1. SKGF Cover Letter;

- 2. Petition For Extension of Time Under 37 C.F.R. § 1.136(a)(1);
- 3. Fee Transmittal Form;
- 4. United States Patent & Trademark Office Credit Card Payment Form;
- 5. Amendment and Reply Under 37 C.F.R. § 1.111;
- 6. Supplemental Request to Approve Proposed Drawing Corrections with 7 sheets of drawings containing proposed corrections marked in red attached;
- 7. Letter to PTO Draftsman: Submission of Formal Drawings;
- 8. Copies of the Formal Drawings (38 sheets); and
- 9. Return postcard.



Please Date Stamp And Return To Our Courier

Appl. No. 09/720,086; 102(e): July 23, 2001
Dkt. No. 0609.4560002/IAG/KRM/DJN; Group Art Unit: 1642
Inventors: Li et al.; Tel: 202/371-2600
Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptide and Uses Thereof

Mouse Dnmt3a DNA sequence

GAATTCCGGC CTGCTGCCGG GCCGCCCGAC CCGCCGGGCC ACACGGCAGA GCCGCCTGAA GCCCAGCGCT GAGGCTGCAC TTTTCCGAGG GCTTGACATC 51 101 AGGGTCTATG TITAAGTCTT AGCTCTTGCT TACAAAGACC ACGGCAATTC CTTCTCTGAA GCCCTCGCAG CCCCACAGCG CCCTCGCAGC CCCAGCCTGC CGCCTACTGC CCAGCAATGC CCTCCAGCGG CCCCGGGGAC ACCAGCAGCT 201 CCTCTCTGGA GCGGGAGGAT GATCGAAAGG AAGGAGAGGA ACAGGAGGAG 301 AACCGTGGCA AGGAAGAGCG CCAGGAGCCC AGCGCCACGG CCCGGAAGGT 351 GGGGAGGCCT GGCCGGAAGC GCAAGCACCC ACCGGTGGAA AGCAGTGACA CCCCCAAGGA CCCAGCAGTG ACCACCAAGT CTCAGCCCAT GGCCCAGGAC TCTGGCCCCT CAGATCTGCT ACCCAATGGA GACTTGGAGA AGCGGAGTGA 451 501 ACCCCAACCT GAGGAGGGGA GCCCAGCTGC AGGGCAGAAG GGTGGGGCCC 551 CAGCTGAAGG AGAGGGAACT GAGACCCCAC CAGAAGCCTC CAGAGCTGTG GAGAATGGCT GCTGTGTGAC CAAGGAAGGC CGTGGAGCCT CTGCAGGAGA 601 651 GGGCAAAGAA CAGAAGCAGA CCAACATCGA ATCCATGAAA ATGGAGGGCT 701 CCCGGGGCCG ACTGCGAGGT GGCTTGGGCT GGGAGTCCAG CCTCCGTCAG CGACCCATGC CAAGACTCAC CTTCCAGGCA GGGGACCCCT ACTACATCAG 801 CAAACGGAAA CGGGATGAGT GGCTGGCACG TTGGAAAAGG GAGGCTGAGA 851 AGAAAGCCAA GGTAATTGCA GTAATGAATG CTGTGGAAGA GAACCAGGCC TCTGGAGAGT CTCAGAAGGT GGAGGAGGCC AGCCCTCCTG CTGTGCAGCA 901 GCCCACGGAC CCTGCTTCTC CGACTGTGGC CACCACCCCT GAGCCAGTAG 1001 GAGGGGATGC TGGGGACAAG AATGCTACCA AAGCAGCCGA CGATGAGCCT 1051 GAGTATGAGG ATGGCCGGGG CTTTGGCATT GGAGAGCTGG TGTGGGGGAA ACTTCGGGGC TTCTCCTGGT GGCCAGGCCG AATTGTGTCT TGGTGGATGA 1101

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Inventors: Liet al.; Tel: 202/371-2600
Title: De Novo DNA Cytosine Methyltransferase Genes, Folypeptides

CAGGCCGGAG CCGAGCAGCT GAAGGCACTC GCTGGGTCAT GTGGTTCGGA 1151 GATGGCAAGT TCTCAGTGGT GTGTGTGGAG AAGCTCATGC CGCTGAGCTC 1201 CTTCTGCAGT GCATTCCACC AGGCCACCTA CAACAAGCAG CCCATGTACC 1251 GCAAAGCCAT CTACGAAGTC CTCCAGGTGG CCAGCAGCCG TGCCGGGAAG 1301 1351 CTGTTTCCAG CTTGCCATGA CAGTGATGAA AGTGACAGTG GCAAGGCTGT 1401 GGAAGTGCAG AACAAGCAGA TGATTGAATG GGCCCTCGGT GGCTTCCAGC CCTCGGGTCC TAAGGGCCTG GAGCCACCAG AAGAAGAGAA GAATCCTTAC 1451 AAGGAAGTTT ACACCGACAT GTGGGTGGAG-CCTGAAGCAG-CTGCTTACGC 1501 CCCACCCCA CCAGCCAAGA AACCCAGAAA GAGCACAACA GAGAAACCTA 1551 AGGTCAAGGA GATCATTGAT GAGCGCACAA GGGAGCGGCT GGTGTATGAG 1601 GTGCGCCAGA AGTGCAGAAA CATCGAGGAC ATTTGTATCT CATGTGGGAG 1651 CCTCAATGTC ACCCTGGAGC ACCCACTCTT CATTGGAGGC ATGTGCCAGA 1701 ACTGTAAGAA CTGCTTCTTG GAGTGTGCTT ACCAGTATGA CGACGATGGG 1751 1801 TACCAGTCCT ATTGCACCAT CTGCTGTGGG GGGCGTGAAG TGCTCATGTG TGGGAACAAC AACTGCTGCA GGTGCTTTTG TGTCGAGTGT GTGGATCTCT 1851 1901 TGGTGGGCC AGGAGCTGCT CAGGCAGCCA TTAAGGAAGA CCCCTGGAAC 1951 TGCTACATGT GCGGGCATAA GGGCACCTAT GGGCTGCTGC GAAGACGGGA 2001 AGACTGGCCT TCTCGACTCC AGATGTTCTT TGCCAATAAC CATGACCAGG AATTTGACCC CCCAAAGGTT TACCCACCTG TGCCAGCTGA GAAGAGGAAG 2051 CCCATCCGCG TGCTGTCTCT CTTTGATGGG ATTGCTACAG GGCTCCTGGT 2101 GCTGAAGGAC CTGGGCATCC AAGTGGACCG CTACATTGCC TCCGAGGTGT 2151 GTGAGGACTC CATCACGGTG GGCATGGTGC GGCACCAGGG AAAGATCATG 2201 2251 TACGTCGGGG ACGTCCGCAG CGTCACACAG AAGCATATCC AGGAGTGGGG CCCATTCGAC CTGGTGATTG GAGGCAGTCC CTGCAATGAC CTCTCCATTG 2301

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This: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

TCAACCCTGC CCGCAAGGGA CTTTATGAGG GTACTGGCCG CCTCTTCTTT 2351 GAGTTCTACC GCCTCCTGCA TGATGCGCGG CCCAAGGAGG GAGATGATCG 2401 CCCCTTCTTC TGGCTCTTTG AGAATGTGGT GGCCATGGGC GTTAGTGACA 2451 AGAGGGACAT CTCGCGATTT CTTGAGTCTA ACCCCGTGAT GATTGACGCC 2501 AAAGAAGTGT CTGCTGCACA CAGGGCCCGT TACTTCTGGG GTAACCTTCC TGGCATGAAC AGGCCTTTGG CATCCACTGT GAATGATAAG CTGGAGCTGC 2601 AAGAGTGTCT GGAGCACGGC AGAATAGCCA AGTTCAGCAA AGTGAGGACC 2651 2701 ATTACCACCA GGTCAAACTC TATAAAGCAG GGCAAAGACC AGCATTTCCC 2751 CGTCTTCATG AACGAGAAGG AGGACATCCT GTGGTGCACT GAAATGGAAA GGGTGTTTGG CTTCCCCGTC CACTACACAG ACGTCTCCAA CATGAGCCGC 2801 TTGGCGAGGC AGAGACTGCT GGGCCGATCG TGGAGCGTGC CGGTCATCCG 2851 2901 CCACCTCTTC GCTCCGCTGA AGGAATATTT TGCTTGTGTG TAAGGGACAT GGGGGCAAAC TGAAGTAGTG ATGATAAAAA AGTTAAACAA ACAAACAAAC 2951 3001 AAAAAACAAA ACAAAACAAT AAAACACCAA GAACGAGAGG ACGGAGAAAA GTTCAGCACC CAGAAGAGAA AAAGGAATTT AAAGCAAACC ACAGAGGAGG 3051 AAAACGCCGG AGGGCTTGGC CTTGCAAAAG GGTTGGACAT CATCTCCTGA 3101 GTTTTCAATG TTAACCTTCA GTCCTATCTA AAAAGCAAAA TAGGCCCCTC 3151 CCCTTCTTCC CCTCCGGTCC TAGGAGGCGA ACTITITGTT TICTACTCTT 3201 TITCAGAGGG GITTICIGIT IGITIGGGIT ITIGITICIT GCIGIGACTG 3251 AAACAAGAGA GTTATTGCAG CAAAATCAGT AACAACAAAA AGTAGAAATG 3301 3351 CCTTGGAGAG GAAAGGGAGA GAGGGAAAAT TCTATAAAAA CTTAAAATAT TGGTTTTTTT TTTTTTCCT TTTCTATATA TCTCTTTGGT TGTCTCTAGC 3401 3451 CTGATCAGAT AGGAGCACAA ACAGGAAGAG AATAGAGACC CTCGGAGGCA GAGTCTCCTC TCCCACCCCC CGAGCAGTCT CAACAGCACC ATTCCTGGTC

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3601 AGACACTTTC TACAGTATTT CAGGTGCCTA CCACACAGGA AACCTTGAAG AAAACCAGTT TCTAGAAGCC GCTGTTACCT_CTTGTTTACA GTTTATATAT 3651 ATATGATAGA TATGAGATAT ATATATAA AAGGTACTGT TAACTACTGT ACATCCCGAC TTCATAATGG TGCTTTCAAA ACAGCGAGAT GAGCAAAGAC 3751 ATCAGCTTCC GCCTGGCCCT CTGTGCAAAG GGTTTCAGCC CAGGATGGGG 3801 3851 AGAGGGGAGC AGCTGGAGGG GGTTTTAACA AACTGAAGGA TGACCCATAT CACCCCCAC CCCTGCCCCA TGCCTAGCTT CACCTGCCAA AAAGGGGCTC 3901 AGCTGAGGTG GTCGGACCCT GGGGAAGCTG AGTGTGGAAT TTATCCAGAC 3951 TCGCGTGCAA TAACCTTAGA ATATGAATCT AAAATGACTG CCTCAGAAAA 4001 4051 ATGGCTTGAG AAAACATTGT CCCTGATTTT GAATTCGTCA GCCACGTTGA 4101 AGGCCCCTTG TGGGATCAGA AATATTCCAG AGTGAGGGAA AGTGACCCGC 4151 CATTAACCCC NCCTGGAGCA AATAAAAAA CATACAAAAT GT

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Mouse Dnmt3b1 DNA Sequence

GAATTCCGGG CGCCGGGGTT AAGCGGCCCA AGTAAACGTA GCGCAGCGAT CGGCGCCGGA GATTCGCGAA CCCGACACTC CGCGCCGCCC GCCGGCCAGG 101 ACCCGCGCG CGATCGCGGC GCCGCGCTAC AGCCAGCCTC ACGACAGGCC CGCTGAGGCT TGTGCCAGAC CTTGGAAACC TCAGGTATAT ACCTTTCCAG 151 201 ACGCGGGATC TCCCCTCCCC CATCCATAGT GCCTTGGGAC CAAATCCAGG 251 GCCTTCTTC AGGAAACAAT GAAGGGAGAC AGCAGACATC TGAATGAAGA AGAGGGTGCC AGCGGGTATG AGGAGTGCAT TATCGTTAAT GGGAACTTCA 301 351 GTGACCAGTC CTCAGACACG AAGGATGCTC CCTCACCCCC AGTCTTGGAG 401 GCAATCTGCA CAGAGCCAGT CTGCACACCA GAGACCAGAG GCCGCAGGTC 451 AAGCTCCCGG CTGTCTAAGA GGGAGGTCTC CAGCCTTCTG AATTACACGC AGGACATGAC AGGAGATGGA GACAGAGATG ATGAAGTAGA TGATGGGAAT 501 GGCTCTGATA TTCTAATGCC AAAGCTCACC CGTGAGACCA AGGACACCAG GACGCGCTCT GAAAGCCCGG CTGTCCGAAC CCGACATAGC AATGGGACCT 601 651 CCAGCTTGGA GAGGCAAAGA GCCTCCCCCA GAATCACCCG AGGTCGGCAG GGCCGCCACC ATGTGCAGGA GTACCCTGTG GAGTTTCCGG CTACCAGGTC 701 751 TCGGAGACGT CGAGCATCGT CTTCAGCAAG CACGCCATGG TCATCCCCTG CCAGCGTCGA CTTCATGGAA GAAGTGACAC CTAAGAGCGT CAGTACCCCA 801 851 TCAGTTGACT TGAGCCAGGA TGGAGATCAG GAGGGTATGG ATACCACACA GGTGGATGCA GAGAGCAGAG ATGGAGACAG CACAGAGTAT CAGGATGATA 901 AAGAGTTTGG AATAGGTGAC CTCGTGTGGG GAAAGATCAA GGGCTTCTCC 951 1001 TGGTGGCCTG CCATGGTGGT GTCCTGGAAA GCCACCTCCA AGCGACAGGC

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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

CATGCCCGGA ATGCGCTGGG TACAGTGGTT TGGTGATGGC AAGTTTTCTG 1051 AGATOTOTIC TGACAAACTG GTGGCTCTGG GGCTGTTCAG CCAGCACTTT 1101 AATCTGGCTA CCTTCAATAA GCTGGTTTCT TATAGGAAGG CCATGTACCA 1151 CACTCTGGAG AAAGCCAGGG TTCGAGCTGG CAAGACCTTC TCCAGCAGTC 1201 CTGGAGAGTC ACTGGAGGAC CAGCTGAAGC CCATGCTGGA GTGGGCCCAC GGTGGCTTCA AGCCTACTGG GATCGAGGGC CTCAAACCCA ACAAGAAGCA 1301 ACCAGTGGTT AATAAGTCGA AGGTGCGTCG TTCAGACAGT AGGAACTTAG 1.351 1401 AACCCAGGAG ACGCGAGAAC AAAAGTCGAA GACGCACAAC°CAATGACTCT GCTGCTTCTG AGTCCCCCCC ACCCAAGCGC CTCAAGACAA ATAGCTATGG 1451 CGGGAAGGAC CGAGGGGAGG ATGAGGAGAG CCGAGAACGG ATGGCTTCTG 1501 AAGTCACCAA CAACAAGGGC AATCTGGAAG ACCGCTGTTT GTCCTGTGGA 1551 AAGAAGAACC CTGTGTCCTT CCACCCCCTC TTTGAGGGTG GGCTCTGTCA 1601 GAGTTGCCGG GATCGCTTCC TAGAGCTCTT CTACATGTAT GATGAGGACG GCTATCAGTC CTACTGCACC GTGTGCTGTG AGGGCCGTGA ACTGCTGCTG 1701 TGCAGTAACA CAAGCTGCTG CAGATGCTTC TGTGTGGAGT GTCTGGAGGT 1751 GCTGGTGGGC GCAGGCACAG CTGAGGATGC CAAGCTGCAG GAACCCTGGA 1801 GCTGCTATAT GTGCCTCCCT CAGCGCTGCC ATGGGGTCCT CCGACGCAGG 1851 1901 AAAGATTGGA ACATGCGCCT GCAAGACTTC TTCACTACTG ATCCTGACCT GGAAGAATTT GAGCCACCCA AGTTGTACCC AGCAATTCCT GCAGCCAAAA 1951 GGAGGCCCAT TAGAGTCCTG TCTCTGTTTG ATGGAATTGC AACGGGGTAC 2001 2051 TTGGTGCTCA AGGAGTTGGG TATTAAAGTG GAAAAGTACA TTGCCTCCGA 2101 AGTCTGTGCA GAGTCCATCG CTGTGGGAAC TGTTAAGCAT GAAGGCCAGA TCAAATATGT CAATGACGTC CGGAAAATCA CCAAGAAAAA TATTGAAGAG 2151 TGGGGCCCGT TCGACTTGGT GATTGGTGGA AGCCCATGCA ATGATCTCTC 2201

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TAACGTCAAT CCTGCCCGCA AAGGTTTATA TGAGGGCACA GGAAGGCTCT 2251 TCTTCGAGTT TTACCACTTG CTGAATTATA CCCGCCCCAA GGAGGGCGAC 2301 AACCGTCCAT TCTTCTGGAT GTTCGAGAAT GTTGTGGCCA TGAAAGTGAA 2351 TGACAAGAAA GACATCTCAA GATTCCTGGC ATGTAACCCA GTGATGATCG 2401 ATGCCATCAA GGTGTCTGCT GCTCACAGGG CCCGGTACTT CTGGGGTAAC 2451 CTACCCGGAA TGAACAGGCC CGTGATGGCT TCAAAGAATG ATAAGCTCGA 2501 GCTGCAGGAC TGCCTGGAGT TCAGTAGGAC AGCAAAGTTA AAGAAAGTGC 2551 AGACAATAAC CACCAAGTCG AACTCCATCA GACAGGGCAA AAACCAGCTT 2601 TTCCCTGTAG TCATGAATGG CAAGGACGAC GTTTTGTGGT GCACTGAGCT 2651 CGAAAGGATC TTCGGCTTCC CTGCTCACTA CACGGACGTG TCCAACATGG 2701 GCCGCGCGC CCGTCAGAAG CTGCTGGGCA GGTCCTGGAG TGTACCGGTC 2751 ATCAGACACC TGTTTGCCCC CTTGAAGGAC TACTTTGCCT GTGAATAGTT 2801 CTACCCAGGA CTGGGGAGCT CTCGGTCAGA GCCAGTGCCC AGAGTCACCC 2851 CTCCCTGAAG GCACCTCACC TGTCCCCTTT TTAGCTCACC TGTGTGGGGC 2901 CTCACATCAC TGTACCTCAG CTTTCTCCTG CTCAGTGGGA GCAGAGCCTC 2951 CTGGCCCTTG CAGGGGAGCC CCGGTGCTCC CTCCGTGTGC ACAGCTCAGA 3001 CCTGGCTGCT TAGAGTAGCC CGGCATGGTG CTCATGTTCT CTTACCCTGA 3051 3101 AACTITAAAA CTIGAAGTAG GTAGTAAGAT GGCTTTCTTT TACCCTCCTG AGTITATCAC TCAGAAGTGA TGGCTAAGAT ACCAAAAAAA CAAACAAAAA CAGAAACAAA AAACAAAAAA AAACCTCAAC AGCTCTCTTA GTACTCAGGT 3201 TCATGCTGCA AAATCACTTG AGATTTTGTT TTTAAGTAAC CCGTGCTCCA CATTIGCTGG AGGATGCTAT TGTGAATGTG GGCTCAGATG AGCAAGGTCA 3301 AGGGGCCAAA AAAAATTCCC CCTCTCCCCC CAGGAGTATT TGAAGATGAT GTTTATGGTT TAAGTCTTCC TGGCACCTTC CCCTTGCTTT GGTACAAGGG

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Title: De Novo DNA Cytosine Methyltransferase Genes, Potypeptides
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CTGAAGTCCT GTTGGTCTTG TAGCATTTCC CAGGATGATG ATGTCAGCAG GGATGACATC ACCACCTTTA GGGCTTTTCC CTGGCAGGGG CCCATGTGGC 3501 3551 TAGTCCTCAC GAAGACTGGA GTAGAATGTT TGGAGCTCAG GAAGGGTGGG 3601 TGGAGTGGCC CTCTTCCAGG TGTGAGGGAT ACGAAGGAGG AAGCTTAGGG 3651 AAATCCATTC CCCACTCCCT CTTGCCAAAT GAGGGGCCCA GTCCCCAACA 3701 GCTCAGGTCC CCAGAACCCC CTAGTTCCTC ATGAGAAGCT AGGACCAGAA 3751 GCACATCGTT CCCCTTATCT GAGCAGTGTT TGGGGAACTA CAGTGAAAAC 3801 CTTCTGGAGA TGTTAAAAGC TTTTTACCCC ACGATAGATT GTGTTTTTAA 3851 GGGGTGCTTT TTTTAGGGGC ATCACTGGAG ATAAGAAAGC TGCATTTCAG 3901 AAATGCCATC GTAATGGTTT TTAAACACCT TTTACCTAAT TACAGGTGCT 3951 ATTITATAGA AGCAGACAAC ACTICTITIT ATGACTCICA GACTICTATI 4001 TICATGTTAC CATTITITIT GTAACTCGCA AGGTGTGGGC TITTGTAACT 4051 TCACAGGTGT GGGGAGAGAC TGCCTTGTTT CAACAGTTTG TCTCCACTGG 4101 TITCTAATTT TTAGGTGCAA AGATGACAGA TGCCCAGAGT TTACCTTTCT

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Human DNMT3A DNA Sequence

1 GCCGCGG CACCAGGGGG CGCAGCCGGG CCGGCCCGAC CCCACCGGCC ATACGGTGGA GCCATCGAAG CCCCCACCCA CAGGCTGACA GAGGCACCGT TCACCAGAGG GCTCAACACC GGGATCTATG TITAAGTITT AACTCTCGCC TCCAAAGACC ACGATAATTC CTTCCCCAAA 128 GCCCAGCAGC CCCCCAGCCC CGCGCAGCCC CAGCCTGCCT CCCGGCGCCCC 178 AGATGCCCGC CATGCCCTCC AGCGGCCCCG GGGACACCAG CAGCTCTGCT 228 GCGGAGCGGG AGGAGGACCG AAAGGACGGA GAGGAGCAGG AGGAGCCGCG 278 328 TGGCAAGGAG GAGCGCCAAG AGCCCAGCAC CACGGCACGG AAGGTGGGGC GGCCTGGGAG GAAGCGCAAG CACCCCCGG TGGAAAGCGG TGACACGCCA 378 AAGGACCCTG CGGTGATCTC CAAGTCCCCA TCCATGGCCC AGGACTCAGG 428 CGCCTCAGAG CTATTACCCA ATGGGGACTT GGAGAAGCGG AGTGAGCCCC 478 AGCCAGAGGA GGGGAGCCCT GCTGGGGGGC AGAAGGGCGG GGCCCCAGCA 528 GAGGGAGAGG GTGCAGCTGA GACCCTGCCT GAAGCCTCAA GAGCAGTGGA 578 628 AAATGGCTGC TGCACCCCCA AGGAGGGCCG AGGAGCCCCT GCAGAAGCGG GCAAAGAACA GAAGGAGACC AACATCGAAT CCATGAAAAT GGAGGGCTCC 678 CGGGGCCGC TGCGGGTGG CTTGGGCTGG GAGTCCAGCC TCCGTCAGCG 728 GCCCATGCCG AGGCTCACCT TCCAGGCGGG GGACCCCTAC TACATCAGCA 778 AGCGCAAGCG GGACGAGTGG CTGGCACGCT GGAAAAGGGA GGCTGAGAAG 828 878 AAAGCCAAGG TCAGTGCAGG AATGAATGCT GTGGAAGAAA ACCAGGGGCC CGGGGAGTCT CAGAAGGTGG AGGAGGCCAG CCCTCCTGCT GTGCAGCAGC 928 CCACTGACCC CGCATCCCCC ACTGTGGCTA CCACGCCTGA GCCCGTGGGG 978 TCCGATGCTG GGGACAAGAA TGCCACCAAA GCAGGCGATG ACGAGCCAGA 1028

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GTACGAGGAC GGCCGGGGT TTGGCATTGG GGAGCTGGTG TGGGGGAAAC TGCGGGGCTT CTCCTGGTGG CCAGGCCGCA TTGTGTCTTG GTGGATGACG GCCCGCAGCC GAGCAGCTGA AGGCACCCGC TGGGTCATGT GGTTCGGAGA CGGCAAATTC TCAGTGGTGT GTGTTGAGAA GCTGATGCCG CTGAGCTCGT TTTGCAGTGC GTTCCACCAG GCCACGTACA ACAAGCAGCC CATGTACCGC AAAGCCATCT ACGAGGTCCT GCAGGTGGCC AGCAGCCGCG CGGGGAAGCT 1328 GTTCCCGGTG TGCCACGACA GCGATGAGAG TGACACTGCC AAGGCCGTCG AGGTGCAGAA CAAGCCCATG ATTGAATGGG CCCTGGGGGG CTTCCAGCCT 1478 AGAAGTGTAC ACGGACATGT GGGTGGAACC TGAGGCAGCT GCCTACGCAC 1528 CACCTCCACC AGCCAAAAAG CCCCGGAAGA GCACAGCGGA GAAGCCCAAG 1578 GTCAAGGAGA TTATTGATGA GCGCACAAGA GAGCGGCTGG TGTACGAGGT 1628 GCGCAGAAG TGCCGGAACA TTGAGGACAT CTGCATCTCC TGTGGGAGCC 1678 1728 TCAATGTTAC CCTGGAACAC CCCCTCTTCG TTGGAGGAAT GTGCCAAAAC 1778 TGCAAGAACT GCTTTCTGGA GTGTGCGTAC CAGTACGACG ACGACGGCTA CCAGTCCTAC TGCACCATCT GCTGTGGGGG CCGTGAGGTG CTCATGTGCG 1828 GAAACAACAA CTGCTGCAGG TGCTTTTGCG TGGAGTGTGT GGACCTCTTG GTGGGGCCGG GGGCTGCCCA GGCAGCCATT AAGGAAGACC CCTGGAACTG 1928 CTACATGTGC GGGCACAAGG GTACCTACGG GCTGCTGCGG CGGCGAGAGG 2028 ACTGGCCCTC CCGGCTCCAG ATGTTCTTCG CTAATAACCA CGACCAGGAA TTTGACCCTC CAAAGGTTTA CCCACCTGTC CCAGCTGAGA AGAGGAAGCC CATCCGGGTG CTGTCTCTCT TTGATGGAAT CGCTACAGGG CTCCTGGTGC 2128 TGAAGGACTT GGGCATTCAG GTGGACCGCT ACATTGCCTC GGAGGTGTGT 2178

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GAGGACTICA TOACGGTGGG CATGGTGCGG CACCAGGGGA AGATCATGTA CGTCGGGGAC GTCCGCAGCG TCACACAGAA GCATATCCAG GAGTGGGGCC 2278 CATTCGATCT GGTGATTGGG GGCAGTCCCT GCAATGACCT CTCCATCGTC 2328 2378 AACCCTGCTC GCAAGGGCCT CTACGAGGGC ACTGGCCGGC TCTTCTTTGA GTTCTACCGC CTCCTGCATG ATGCGCGGCC CAAGGAGGGA GATGATCGCC 2428 CCTTCTTCTG GCTCTTTGAG AATGTGGTGG CCATGGGCGT TAGTGACAAG AGGGACATCT CGCGATTTCT CGAGTCCAAC CCTGTGATGA TTGATGCCAA 2528 AGAAGTGTCA GCTGCACACA GGGCCCGCTA CTTCTGGGGT AACCTTCCCG 2578 GTATGAACAG GCCGTTGGCA TCCACTGTGA ATGATAAGCT GGAGCTGCAG 2628 GAGTGTCTGG AGCATGGCAG GATAGCCAAG TTCAGCAAAG TGAGGACCAT TACTACGAGG TCAAACTCCA TAAAGCAGGG CAAAGACCAG CATTTTCCTG 2728 TCTTCATGAA TGAGAAAGAG GACATCTTAT GGTGCACTGA AATGGAAAGG 2778 GTATTTGGTT TCCCAGTCCA CTATACTGAC GTCTCCAACA TGAGCCGCTT 2828 GGCGAGGCAG AGACTGCTGG GCCGGTCATG GAGCGTGCCA GTCATCCGCC 2878 ACCTCTTCGC TCCGCTGAAG GAGTATTTTG CGTGTGTGTA AGGGACATGG 2928 2978 GGGCAAACTG AGGTAGCGAC ACAAAGTTAA ACAAACAAAC AAAAAAACACA 3028 AAACATAATA AAACACCAAG AACATGAGGA TGGAGAGAAG TATCAGCACC CAGAAGAGAA AAAGGAATTT AAAACAAAAA CCACAGAGGC GGAAATACCG 3078 3128 GAGGGCTTTG CCTTGCGAAA AGGGTTGGAC ATCATCTCCT GATTTTTCAA 3178 TCCCCCTTCC CTTTTTTTC GGTCAGACCT TTTATTTTCT ACTCTTTCA 3228 GAGGGGTTTT CTGTTTGTTT GGGTTTTGTT TCTTGCTGTG ACTGAAACAA 3278 GAAGGTTATT GCAGCAAAAA TCAGTAACAA AAAATAGTAA CAATACCTTG 3328

CAGAGGAAAG GTGGGAGGAG AGGAAAAAAG GGAAATTTTT AAAGAAATCT

3378

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3428 ATATATIGGG TIGITITIT TITIGITITI TGTTTTTTT TTTTGGGTTT TITTTTTTTA CTATATATCT TTTTTTTGTT GTCTCTAGCC TGATCAGATA 3478 GGAGCACAAG CAGGGGACGG AAAGAGAGAG ACACTCAGGC GGCAGCATTC 3528 CCTCCCAGCC ACTGAGCTGT CGTGCCAGCA CCATTCCTGG TCACGCAAAA 3628 CAGAACCCAG TTAGCAGCAG GGAGACGAGA ACACCACACA AGACATTTT 3678 CTACAGTATT TCAGGTGCCT ACCACACAGG AAACCTTGAA GAAAATCAGT 3728 TTCTAGAAGC CGCTGTTACC TCTTGTTTAC AGTTTATATA TATATGATAG ATATGAGATA TATATAAAA AGGTACTGTT AACTACTGTA CAACCCGACT 3778 3828 TCATAATGGT GCTTTCAAAC AGCGAGATGA GTAAAAACAT CAGCTTCCAC 3878 GTTGCCTTCT GCGCAAAGGG TTTCACCAAG GATGGAGAAA GGGAGACAGC 3928 TTGCAGATGG CGCGTTCTCA CGGTGGGCTC TTCCCCTTGG TTTGTAACGA AGTGAAGGAG GAGAACTTGG GAGCCAGGTT CTCCCTGCCA AAAAGGGGGC 3978 4028 TAGATGAGGT GGTCGGGCCC GTGGACAGCT GAGAGTGGGA TTCATCCAGA CTCATGCAAT AACCCTTTGA TTGTTTTCTA AAAGGAGACT CCCTCGGCAA 4078 GATGGCAGAG GGTACGGAGT CTTCAGGCCC AGTTTCTCAC TTTAGCCAAT 4128 4178 TCGAGGGCTC CTTGTGGTGG GATCAGAACT AATCCAGAGT GTGGGAAAGT GACAGTCAAA ACCCCACCTG GAGCAAATAA AAAAACATAC AAAACGTAAA 4278 AAAAAAAAA AAAAAA

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Title: De Novo DNA Cytosins Methyltransferase Genes, Polypeptides and Uses Thereof

Human DNMT3B1 DNA Sequence:

GGCCGCGAAT TCGGCACGAG CCCTGCACGG CCGCCAGCCG GCCTCCCGCC AGCCAGCCC GACCCGCGC TCCGCCGCCC AGCCGCCC CAGCCAGCCC 101 TGCGGCAGGA AAGCATGAAG GGAGACACCA GGCATCTCAA TGGAGAGGAG GACGCCGGCG GGAGGGAAGA CTCGATCCTC GTCAACGGGG CCTGCAGCGA 201 CCAGTCCTCC GACTCGCCCC CAATCCTGGA GGCTATCCGC ACCCCGGAGA 251 TCAGAGGCCG AAGATCAAGC TCGCGACTCT CCAAGAGGGA GGTGTCCAGT CTGCTAAGCT ACACACAGGA CTTGACAGGC GATGGCGACG GGGAAGATGG 351 GGATGGCTCT GACACCCCAG TCATGCCAAA GCTCTTCCGG GAAACCAGGA CTCGTTCAGA AAGCCCAGCT GTCCGAACTC GAAATAACAA CAGTGTCTCC 451 AGCCGGGAGA GGCACAGGCC TTCCCCACGT TCCACCCGAG GCCGGCAGGG 501 CCGCAACCAT GTGGACGAGT CCCCCGTGGA GTTCCCGGCT ACCAGGTCCC 551 TGAGACGCC GCCAACAGCA TCGGCAGGAA CGCCATGGCC GTCCCCTCCC 601 AGCTCTTACC TTACCATCGA CCTCACAGAC GACACAGAGG ACACACATGG 651 GACGCCCCAG AGCAGCAGTA CCCCCTACGC CCGCCTAGCC CAGGACAGCC AGCAGGGGG CATGGAGTCC CCGCAGGTGG AGGCAGACAG TGGAGATGGA 751 GACACTTCAG AGTATCAGGA TGGGAAGGAG TTTGGAATAG GGGACCTCGT 801 GTGGGGAAAG ATCAAGGGCT TCTCCTGGTG GCCCGCCATG GTGGTGTCTT 851 GGAAGGCCAC CTCCAAGCGA CAGGCTATGT CTGGCATGCG GTGGGTCCAG 901 TGGTTTGGCG ATGGCAAGTT CTCCGAGGTC TCTGCAGACA AACTGGTGGC 951 ACTGGGGCTG TTCAGCCAGC ACTTTAATTT GGCCACCTTC AATAAGCTCG 1001 TCTCCTATCG AAAAGCCATG TACCATGCTC TGGAGAAAGC TAGGGTGCGA 1051 GCTGGCAAGA CCTTCCCCAG CAGCCCTGGA GACTCATTGG AGGACCAGCT GAAGCCCATG TTGGAGTGGG CCCACGGGGG CTTCAAGCCC ACTGGGATCG 1101 AGGGCCTCAA ACCCAACAAC ACGCAACCAG TGGTTAATAA GTCGAAGGTG

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Dkt. No. 0609.4560002/JAG/KRM/DJN; Group Art Unit: 1642
Inventor: Li et al.; Tel: 202/371-2600
Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

CGTCGTGCAG GCAGTAGGAA ATTAGAATCA AGGAAATACG AGAACAAGAC 1201 1251 TCGAAGACGC ACAGCTGACG ACTCAGCCAC CTCTGACTAC TGCCCCGCAC 1301 CCAAGCGCCT CAAGACAAAT TGCTATAACA ACGGCAAAGA CCGAGGGGAT GAAGATCAGA GCCGAGAACA AATGGCTTCA GATGTTGCCA ACAACAAGAG 1351 CAGCCTGGAA GATGGCTGTT TGTCTTGTGG CAGGAAAAAC CCCGTGTCCT 1401 TCCACCCTCT CTTTGAGGGG GGGCTCTGTC AGACATGCCG GGATCGCTTC 1451 1501 CTTGAGCTGT TTTACATGTA TGATGACGAT GGCTATCAGT CTTACTGCAC 1551 TGTGTGCTGC GAGGGCCGAG AGCTGCTGCT TTGCAGCAAC ACGAGCTGCT GCCGGTGTTT CTGTGTGGAG TGCCTGGAGG TGCTGGTGGG CACAGGCACA 1601 1651 GCGCCGAGG CCAAGCTTCA GGAGCCCTGG AGCTGCTACA TGTGTCTCCC 1701 GCAGCGCTGT CATGGCGTCC TGCGGCGCCG GAAGGACTGG AACGTGCGCC 1751 TGCAGGCCTT CTTCACCAGT GACACGGGGC TTGAATACGA AGCCCCCAAG CTGTACCCTG CCATTCCCGC AGCCCGAAGG CGGCCCATTC GAGTCCTGTC 1801 ATTGTTTGAT GGCATCGCGA CAGGCTACCT AGTCCTCAAA GAGTTGGGCA 1851 TAAAGGTAGG AAAGTACGTC GCTTCTGAAG TGTGTGAGGA GTCCATTGCT 1901 1951 GTTGGAACCG TGAAGCACGA GGGGAATATC AAATACGTGA ACGACGTGAG GAACATCACA AAGAAAAATA TTGAAGAATG GGGCCCATTT GACTTGGTGA 2001 2051 TTGGCGGAAG CCCATGCAAC GATCTCTCAA ATGTGAATCC AGCCAGGAAA GGCCTGTATG AGGGTACAGG CCGGCTCTTC TTCGAATTTT ACCACCTGCT 2101 2151 GAATTACTCA CGCCCCAAGG AGGGTGATGA CCGGCCGTTC TTCTGGATGT 2201 TTGAGAATGT TGTAGCCATG AAGGTTGGCG ACAAGAGGGA CATCTCACGG 2251 TTCCTGGAGT GTAATCCAGT GATGATTGAT GCCATCAAAG TTTCTGCTGC 2301 TCACAGGCC CGATACTTCT GGGGCAACCT ACCCGGGATG AACAGGCCCG 2351 TGATAGCATC AAAGAATGAT AAACTCGAGC TGCAGGACTG CTTGGAATAC AATAGGATAG CCAAGTTAAA GAAAGTACAG ACAATAACCA CCAAGTCGAA

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CTCGATCAAA CAGGGGAAAA ACCAACTTTT CCCTGTTGTC ATGAATGGCA 2451 2501 AAGAAGATGT TTTGTGGTGC ACTGAGCTCG AAAGGATCTT TGGCTTTCCT GTGCACTACA CAGACGTGTC CAACATGGGC CGTGGTGCCC GCCAGAAGCT 2551 GCTGGGAAGG TCCTGGAGCG TGCCTGTCAT CCGACACCTC TTCGCCCCTC 2601 2651 TGAAGGACTA CTTTGCATGT GAATAGTTCC AGCCAGGCCC CAAGCCCACT GGGGTGTGTG GCAGAGCCAG GACCCAGGAG GTGTGATTCC TGAAGGCATC *2701* 2751 CCCAGGCCCT GCTCTTCCTC AGCTGTGTGG GTCATACCGT GTACCTCAGT TCCCTCTTGC TCAGTGGGGG CAGAGCCACC TGACTCTTGC AGGGGTAGCC, 2801 2851 TGAGGTGCCG CCTCCTTGTG CACAAATCAG ACCTGGCTGC TTGGAGCAGC 2901 CTAACACGGT GCTCATTTTT TCTTCTCCTA AAACTTTAAA ACTTGAAGTA *2*951 GGTAGCAACG TGGCTTTTTT TTTTTCCCTT CCTGGGTCTA CCACTCAGAG AAACAATGGC TAAGATACCA AAACCACAGT GCCGACAGCT CTCCAATACT 3001 CAGGTTAATG CTGAAAAATC ATCCAAGACA GTTATTGCAA GAGTTTAATT 3051 3101 TTTGAAAACT GGGTACTGCT ATGTGTTTAC AGACGTGTGC AGTTGTAGGC 3151 ATGTAGCTAC AGGACATTTT TAAGGGCCCA GGATCGTTTT TTCCCAGGGC AAGCAGAAGA GAAAATGTTG TATATGTCTT TTACCCGGCA CATTCCCCTT 3201 3251 GCCTAAATAC AAGGGCTGGA GTCTGCACGG GACCTATTAG AGTATTTTCC 3301 ACAATGATGA TGATTTCAGC AGGGATGACG TCATCATCAC ATTCAGGGCT 3351 ATTTTTCCC CCACAAACCC AAGGGCAGGG GCCACTCTTA GCTAAATCCC 3401 TCCCCGTGAC TGCAATAGAA CCCTCTGGGG AGCTCAGGAA GGGGTGTGCT 3451 GAGTTCTATA ATATAAGCTG CCATATATTT TGTAGACAAG TATGGCTCCT 3501 CCATATCTCC CTCTTCCCTA GGAGAGGAGT GTGAAGCAAG GAGCTTAGAT 3551 AAGACACCCC CTCAAACCCA TTCCCTCTCC AGGAGACCTA CCCTCCACAG GCACAGGTCC CCAGATGAGA AGTCTGCTAC CCTCATTTCT CATCTTTTA CTAAACTCAG AGGCAGTGAC AGCAGTCAGG GACAGACATA CATTTCTCAT 3651

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3701 ACCTTCCCCA CATCTGAGAG ATGACAGGGA AAACTGCAAA GCTCGGTGCT 3751 CCCTTTGGAG ATTITITAAT CCTTTTTTAT TCCATAAGAA GTCGTTTTTA 3801 GGGAGAACGG GAATTCAGAC AAGCTGCATT TCAGAAATGC TGTCATAATG 3851 GTITTTAACA CCTTTTACTC TTCTTACTGG TGCTATTTTG TAGAATAAGG 3901 AACAACGTTG ACAAGTTTTG TGGGGCTTTT TATACACTTT TTAAAATCTC 3951 AAACTICTAT TITTATGTTT AACGTTTTCA TTAAAATTTT TITGTAACTG 4001 GAGCCACGAC GTAACAAATA TGGGGAAAAA ACTGTGCCTT GTTTCAACAG 4051 TITITGCTAA TITITAGGCT GAAAGATGAC GGATGCCTAG AGTTTACCTT 4101 ATGTTTAATT AAAATCAGTA TTTGTCTAAA AAAAAAAAA AAAAA

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Mouse Dnmt3a Protein

MPSSGPGDTS SSSLEREDDR KEGEEQEENR GKEERQEPSA TARKVGRPGR KRKHPPVESS DTPKDPAVTT KSQPMAQDSG PSDLLPNGDL EKRSEPQPEE GSPAAGOKGG APAEGEGTET PPEASRAVEN GCCVTKEGRG ASAGEGKEQK 101 QTNIESMKME GSRGRLRGGL GWESSLRQRP MPRLTFQAGD PYYISKRKRD 151 EWLARWKREA EKKAKVIAVM NAVEENQASG ESQKVEEASP PAVQQPTDPA 201 SPTVATTPEP VGGDAGDKNA TKAADDEPEY EDGRGFGIGE LVWGKLRGFS 251 WWPGRIVSWW MTGRSRAAEG TRWVMWFGDG KFSVVCVEKL MPLSSFCSAF 301 HQATYNKQPM YRKAIYEVLQ VASSRAGKLF PACHDSDESD SGKAVEVQNK 351 OMIEWALGGF OPSCPKGLEP PEEEKNPYKE VYTDMWVEPE AAAYAPPPPA 401 451 KKPRKSTTEK PKVKEIIDER TRERLVYEVR QKCRNIEDIC ISCGSLNVTL EHPLFIGGMC QNCKNCFLEC AYQYDDDGYQ SYCTICCGGR EVLMCGNNNC 501 CRCFCVECVD LLVGPGAAQA AIKEDPWNCY MCGHKGTYGL LRRREDWPSR 551 LOMFFANNHD QEFDPPKVYP PVPAEKRKPI RVLSLFDGIA TGLLVLKDLG 601 IOVDRYIASE VCEDSITVGM VRHQGKIMYV GDVRSVTQKH IQEWGPFDLV 651 IGGSPCNDLS IVNPARKGLY EGTGRLFFEF YRLLHDARPK EGDDRPFFWL 701 FENVVAMOVS DKRDISRFLE SNPVMIDAKE VSAAHRARYF WONLPGMNRP 751 LASTVNDKLE LQECLEHGRI AKFSKVRTIT TRSNSIKQGK DQHFPVFMNE 801 KEDILWCTEM ERVFGFPVHY TDVSNMSRLA RQRLLGRSWS VPVIRHLFAP 851 LKEYFACV* 901

FIG. 2A

Mouse Dnmt3b1 Protein

MKGDSRHLNE EEGASGYEEC IIVNGNFSDQ SSDTKDAPSP PVLEAICTEP 51 VCTPETRGRR SSSRLSKREV SSLLNYTQDM TGDGDRDDEV DDGNGSDILM 101 PKLTRETKDT RTRSESPAVR TRHSNGTSSL ERQRASPRIT RGRQGRHHVQ EYPVEFPATR SRRRRASSSA STPWSSPASV DFMEEVTPKS VSTPSVDLSQ 201 DGDQEGMDTT QVDAESRDGD STEYQDDKEF GIGDLVWGKI KGFSWWPAMV 251 VSWKATSKRQ AMPGMRWVQW FGDGKFSEIS ADKLVALGLF SQHFNLATFN 301 KLVSYRKAMY HTLEKARVRA GKTFSSSPGE SLEDQLKPML EWAHGGFKPT 351 GIEGLKPNKK QPVVNKSKVR RSDSRNLEPR RRENKSRRRT TNDSAASESP PPKRLKTNSY GGKDRGEDEE SRERMASEVT NNKGNLEDRC LSCGKKNPVS 401 FHPLFEGGLC QSCRDRFLEL FYMYDEDGYQ SYCTVCCEGR ELLLCSNTSC CRCFCVECLE VLVGAGTAED AKLQEPWSCY MCLPQRCHGV LRRRKDWNMR 501 LODFFTTDPD LEEFEPPKLY PAIPAAKRRP IRVLSLFDGI ATGYLVLKEL 551 601 GIKVEKYIAS EVCAESIAVG TVKHEGQIKY VNDVRKITKK NIEEWGPFDL 651 VIGGSPCNDL SNVNPARKGL YEGTGRLFFE FYHLLNYTRP KEGDNRPFFW 701 MFENVVAMKV NDKKDISRFL ACNPVMIDAI KVSAAHRARY FWGNLPGMNR 751 PVMASKNDKL ELQDCLEFSR TAKLKKVQTI TTKSNSIRQG KNQLFPVVMN 801 GKDDVLWCTE LERIFGFPAH YTDVSNMGRG ARQKLLGRSW SVPVIRHLFA 851 PLKDYFACE*

FIG. 2B

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Title: De Novo DNA Cytosine Methyliransferase Genes, Polypeptides

Human DNMT3A Protein

MPAMPSSGPG DTSSSAAERE EDRKDGEEGE EPRGKEERGE PSTTARKVGR PGRKRKHPPV ESGDTPKDPA VISKSPSMAQ DSGASELLPN GDLEKRSEPQ 51 PEEGSPAGGO KGGAPAEGEG AAETLPEASR AVENGCCTPK EGRGAPAEAG 101 KEOKETNIES MKMEGSRGRL RGGLGWESSL RQRPMPRLTF QAGDPYYISK 151 RKRDEWLARW KREAEKKAKV JAGMNAVEEN QCPGESQKVE EASPPAVQQP 201 251 TDPASPTVAT TPEPVGSDAG DKNATKAGDD EPEYEDGRGF GIGELVWGKL RGFSWWPGRI VSWWMTGRSR AAEGTRWVMW FGDGKFSVVC VEKLMPLSSF 301 CSAFHOATYN KOPMYRKAIY EVLQVASSRA GKLFPVCHDS DESDTAKAVE 351 VONKPMIEWA LGGFQPSGPK GLEPPEEEKN PYKEVYTDMW VEPEAAAYAP 401 PPPAKKPRKS TAEKPKVKE! IDERTRERLV YEVRQKCRN! EDICISCGSL 451 NVTLEHPLFV GGMCQNCKNC FLECAYQYDD DGYQSYCTIC CGGREVLMCG 501 NNNCCRCFCV ECVDLLVGPG AAQAAIKEDP WNCYMCGHKG TYGLLRRRED 551 WPSRLQMFFA NNHDQEFDPP KVYPPVPAEK RKPIRVLSLF DGIATGLLVL 601 KDLGIQVDRY JASEVCEDSI TVGMVRHQGK IMYVGDVRSV TQKHIQEWGP 651 FDLVIGGSPC NDLSIVNPAR KGLYEGTGRL FFEFYRLLHD ARPKEGDDRP 701 FFWLFENVVA MGVSDKRDIS RFLESNPVMI DAKEVSAAHR ARYFWGNLPG 751 MNRPLASTVN DKLELQECLE HGRIAKFSKV RTITTRSNSI KQGKDQHFPV 801 FMNEKEDILW CTEMERVFGF PVHYTDVSNM SRLARQRLLG RSWSVPVIRH 901 LFAPLKEYFA CV*

FIG. 2C

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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

Human DNMT3B1 Protein

MKGDTRHLNG EEDAGGREDS ILVNGACSDQ SSDSPPILEA IRTPEIRGRR SSSRLSKREV SSLLSYTOOL TODGDGEDGD GSDTPVMPKL FRETRTRSES PAVRTRNNNS VSSRERHRPS PRSTRGRQGR NHVDESPVEF PATRSLRRRA 151 TASACTPWPS PPSSYLTIDL TODTEDTHGT PQSSSTPYAR LAQDSQQGM *2*01 ESPQVEADSG DGDSSEYQDG KEFGIGDLVW GKIKGFSWWP ANVVSWKATS 251 KRQAMSGMRW VQWFGDGKFS EVSADKLVAL GLFSQHFNLA TFNKLVSYRK 301 AMYHALEKAR VRAGKTFPSS PGDSLEDQLK PMLEWAHGGF KPTGIEGLKP 351 NNTQPVVNKS KVRRAGSRKL ESRKYENKTR RRTADDSATS DYCPAPKRLK 401 TNCYNNGKDR GDEDQSREQM ASDVANNKSS LEDGCLSCGR KNPVSFHPLF 451 EGGLCQTCRD RFLELFYMYD DDGYQSYCTV CCEGRELLLC SNTSCCRCFC 501 VECLEVLYGT GTAAEAKLQE PWSCYMCLPQ RCHGVLRRRK DWNVRLQAFF 551 TSDTGLEYEA PKLYPAIPAA RRRPIRVLSL FDGIATGYLV LKELGIKVGK 601 YVASEVCEES IAVGTVKHEG NIKYVNDVRN ITKKNIEEWG PFDLVIGGSP 651 CNDLSNVNPA RKGLYEGTGR LFFEFYHLLN YSRPKEGDDR PFFWMFENVV 701 AMKVGDKRDI SRFLECNPVM IDAIKVSAAH RARYFWGNLP GMNRPVIASK 751 NDKLELQDCL EYNRIAKLKK VQTITTKSNS IKQGKNQLFP VVMNGKEDVL 801 WCTELERIFG FPVHYTDVSN MGRGARQKLL GRSWSVPVIR HLFAPLKDYF 851 ACE*

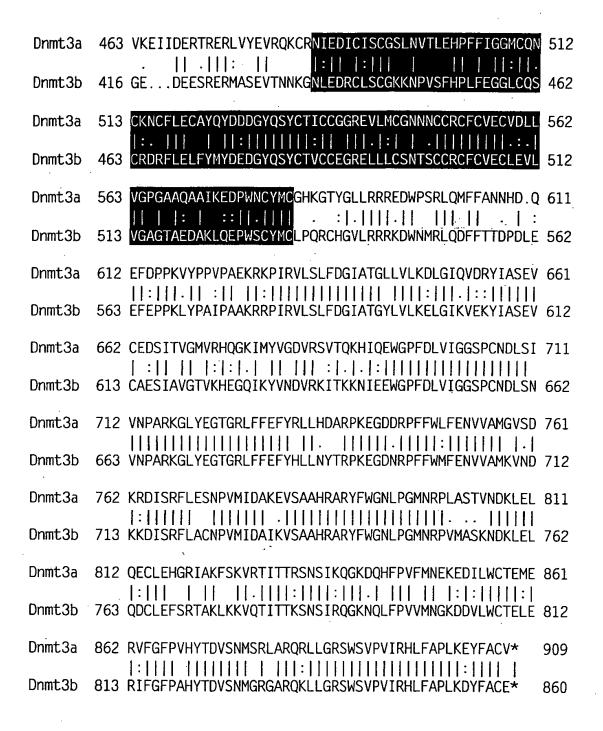
FIG. 2D

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Dkt. No. 0609.4560002/JAG/KRM/DJN; Group Art Unit: 1642
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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

Dnmt3a	1	${\tt MPSSGPGDTSSSSLEREDDRKEGEEQEENRGKEERQEPSATARKVGRPGR}$	50	
Dnmt3a		KRKHPPVESSDTPKDPAVTTKSQPMAQDSGPSDLLPNGDLEKRSEP . . : : . :		
Dnmt3b	1	. . : : . : MKGDSRHLNEEEGASGYEECIIVNGNFSDQSSD	33	
Dnmt3a	97	QPEEGSPAAGQKGGAPAEGEGTETPPEAS.RAVENGCCVTKEGR	139	
Dnmt3b	34	: TKDAPSPPVLEAICTEPVCTPETRGRRSSSRLSKREVSSLLNYTQDMTGD	83	
Dnmt3a	140	GASAGEGKEQKQTNIESMKMEGSRGRLRGGLGWESSLRQ	178	
Dnmt3b	84	:: : GDRDDEVDDGNGSDILMPKLTRETKDTRTRSESPAVRTRHSNGTSSLERQ	133	
Dnmt3a	179	RPMPRLTFQAGDPYYISKRKRDEWLARWKREAEKKAKVIAVMNAVEENQA	228	
Dnmt3b	134	: ::: : ::: RASPRITRGRQGRHHVQEYPVEFPATRSRRRASSSASTPWSSPA	178	
Dnmt3a	229	SGESQKVEEASPPAVQQPTDPASPTVATTPEPVGGDAGDKNATKAADDEP	278	
Dnmt3b	179	: SVDLSQDGDQEGMDTTQVDAESRDGDST	222	
Dnmt3a	279	EYEDGRGFGIGELVWGKLRGFSWWPGRIVSWWMTGRSRAAEGTRWVMWFG	328	
Dnmt3b	223	EYQDDKEFGIGDLVWGKIKGFSWWPAMVVSWKATSKRQAMPGMRWVQWFG	272	
Dnmt3a	329	DGKFSVVCVEKLMPLSSFCSAFHQATYNKQPMYRKAIYEVLQVASSRAGK	378	
Dnmt3b	273	: :	322	
Dnmt3a	379	LFPACHDSDESDSGKAVEVQNKQMIEWALGGFQPSGPKGLEPPEEEKN	426	
Dnmt3b	323	TFSSSPGESLEDQLKPMLEWAHGGFKPTGIEGLKPNKKQPVVN	365	
Dnmt3a	427	PYKEVYTDMW.VEPEAAAYAPPPPAKKPRKSTTEKPK	462	
Dnmt3b	366	. . : : . : : KSKVRRSDSRNLEPRRRENKSRRRTTNDSAASESPPPKRLKTNSYGGKDR	415	

FIG.3A-1

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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

DNMT3A	1 MPAMPSSGPGDTSSSAAEREEDRKDGEEQEEPRGKEERQEPSTTARKVGR
DNMT3A	51 PGRKRKHPPVESGDTPKDPAVISKSPSMAQDSGASELLPNGDLEKRSEPQ
DNMT3B	: 1MKGDTRHLNGEEDAGGREDSILVNGACSDQSSDSP
DNMT3A	101 PEEGSPAGGQKGGAPAEGEGAAETLPEASRAVENGCCTPKEGRGAPAEAG
DNMT3B	. : :. .: 36 PILEAIRTPEIRGGWASSRLSKREVSSLLSYTQDLTGDGDGEDGDGSDTP
DNMT3A	151 KEQKETNIESMKMEGSRGRLRGGLGWESSLRQRPMPRLTFQAGDPYYISK
DNMT3B	86 VMPKLFRETRTRSESPAVRTRNNNSVSSRERHRPSPRSTRGRQGRNHVDE
DNMT3A	201 RKRDEWLARWKREAEKKAKVIAGMNAVEENQGPGESQKVEEASPPAVQQP
DNMT3B	: : 136 SPVEFPATRSLRRRATASAGTPWPSPPSSYLTIDLTDDTEDTHGTPQS
DNMT3A	251 TDPASPTVATTPEPVGSDAGDKNATKAGDDEPEYEDGRGFGIGELVWGKL
DNMT3B	184 SSTPYARLAQDSQQGGMESPQVEADSGDGDSSEYQDGKEFGIGDLVWGKI
DNMT3A	301 RGFSWWPGRIVSWWMTGRSRAAEGTRWVMWFGDGKFSVVCVEKLMPLSSF
DNMT3B	234 KGFSWWPAMVVSWKATSKRQAMSGMRWVQWFGDGKFSEVSADKLVALGLF
DNMT3A	351 CSAFHQATYNKQPMYRKAIYEVLQVASSRAGKLFPVCHDSDESDTAKAVE
DNMT3B	: :
DNMT3A	401 VQNKPMIEWALGGFQPSGPKGLEPPEEEKNPYKEVYTDMWVE
DNMT3B	327 DQLKPMLEWAHGGFKPTGIEGLKPNNTQPVVNKSKVRRAGSRKLESRKYE
DNMT3A	443PEAAAYAPPPPAKKPRKSTAEKPKVKEIIDERTRERLVYEVRQ : . . . :: .: :
DNMT3B	377 NKTRRTADDSATSDYCPAPKRLKTNCYNNGKDRGDEDQSREQMASDVAN

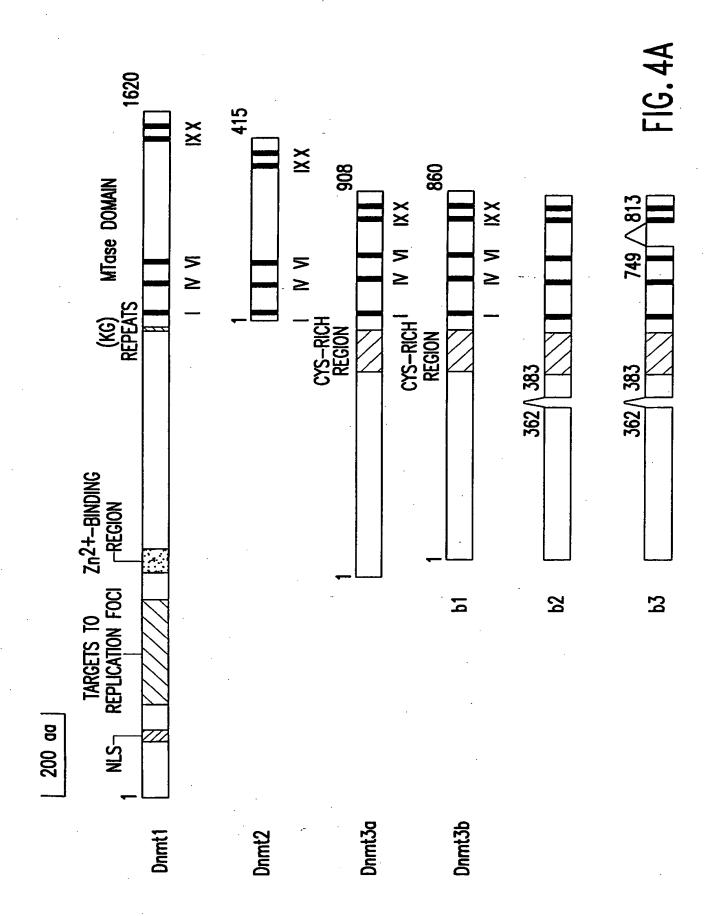
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Inventor: Li et al.; Tel: 202/371-2660
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and Uses Thereof

DNMT3A	486	KCRNIEDICISCGSLNVTLEHPLFVGGMCQNCKNCFLECAYQYDDDGYQS
DNMT3B	427	.: :
DNMT3A	536	YCTICCGGREVLMCGNNNCCRCFCVECVDLLVGPGAAQAAIKEDPWNCYM
DNMT3B	477	YCTVCCEGRELLLCSNTSCCRCFCVECLEVLVGTGTAAEAKLQEPWSCYM
DNMT3A	586	CGHKGTYGLLRRREDWPSRLQMFFANNHDQEFDPPKVYPPVPAEKRKPIR
DNMT3B	527	CLPQRCHGVLRRRKDWNVRLQAFFTSDTGLEYEAPKLYPAIPAARRRPIR
DNMT3A	636	VLSLFDGIATGLLVLKDLGIQVDRYIASEVCEDSITVGMVRHQGKIMYVG
DNMT3B	577	VLSLFDGIATGYLVLKELGIKVGKYVASEVCEESIAVGTVKHEGNIKYVN
DNMT3A	686	DVRSVTQKHIQEWGPFDLVIGGSPCNDLSIVNPARKGLYEGTGRLFFEFY
DNMT3B	627	DVRNITKKNIEEWGPFDLVIGGSPCNDLSNVNPARKGLYEGTGRLFFEFY
DNMT3A	736	RLLHDARPKEGDDRPFFWLFENVVAMGVSDKRDISRFLESNPVMIDAKEV
DNMT3B	677	HLLNYSRPKEGDDRPFFWMFENVVAMKVGDKRDISRFLECNPVMIDAIKV
DNMT3A	786	SAAHRARYFWGNLPGMNRPLASTVNDKLELQECLEHGRIAKFSKVRTITT
DNMT3B	727	SAAHRARYFWGNLPGMNRPVIASKNDKLELQDCLEYNRIAKLKKVQTITT
DNMT3A	836	RSNSIKQGKDQHFPVFMNEKEDILWCTEMERVFGFPVHYTDVSNMSRLAR:
DNMT3B .	777	KSNSIKQGKNQLFPVVMNGKEDVLWCTELERIFGFPVHYTDVSNMGRGAR
DNMT3A	886	QRLLGRSWSVPVIRHLFAPLKEYFACV*
DNMT3B	827	QKLLGRSWSVPVIRHLFAPLKDYFACE*

FIG.3B-2

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Inventors: Li et al.; Tel: 202/371-2600
Täle: De Novo DNA Cytosino Methyltransferase Genes, Polype_{pri}de and Uses Thereof

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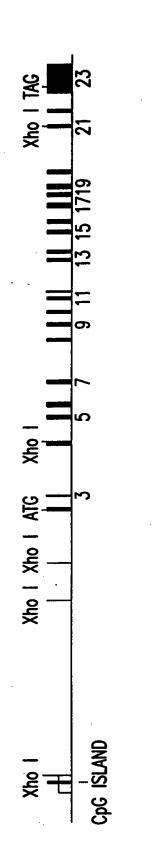
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DNMT3A DNMT3B Zmt3 FIG. 4B

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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof



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17618bp)	887b <u>p</u>)	3343bp)	$1642\overline{bp}$	602bp)	$1403\overline{\text{bp}}$	2588bp)	917bp)	765bp)	1813bp)	115bp)	1095bp)	417bp)	1160bp)	(dq009	824bp)	536bp)	352bp)	958b _D)	2867bb)	801bb)	1434 bp	(A)
.intron(.intron(.intron(.intron(.intron(.intron(.intron(.intron(.intron(.intron(.intron(.intron(.intron(intron(.intron(.intron(.intron(.intron(.intron(.intron(.intron(.intron(
(>=90bp) CGGCAGgtgagggccccggggg.intron(17618bp)	148bp) TCAGAGgtggctgggcagtgg.intron	62bp) ACACAGgtatggtctctgctc.intron	102bp) CCAGCTgtaagtagccacacc.intron	125bp) ACCAGGgttgttccccagatg.intron	22bp) TATCAGgtatggccgagaggg.intron	TCCGAGgtgagtccgggggaag.intron	CTGGAGgtaacatgggatgag.intron	145bp) AACCAGgtgggaatgagtccc.intron	60bp) AATACGgtatttccttcctgt.intron	126bp)GCCGAGgtgattgttggggtac.intron	TGGAAGgtaacgttctctccc.intron	TGCCGGgtaagtcctcctact.intron	CTGCCGgtgagcactgggccc.intron	184bp) GAATACgtaagccacaggctc.intron	85bp) CGACAGgtgagttcgggggaac.intron	146bp) AAAAATgtgagggcagtctgt.intron	TGTATGgtgagcatccttctc.intron	149bp) CTGGAGgtgagggaatctggg.intron	86bp) GAACAGgtaacaaaqqqctct.intron	70bp) GCCAAGttaaagaaagtacag.intron	CGAAAGGtgagcaaggctgca.intron	
) (dq06=<	148bp)	(42pb) 1	102bp)(125bp) A			108bp) (145bp) 7	(dq09	126bp)(113bp)(184bp)(85bp) (146bp) 7	91bp)7	149bp) (86bp) (70bp) (119bp)(_
	Exon2 (Exon3 (Exon4 (Exon5 (Exone (Exon7 (Exon8 (Exon9 (Exon10 (Exon11 (Exon12 (Exon13 (Exon14 (Exon15 (Exon16 (Exon17 (Exon18 (Exon19 (Exon20 (Exon21 (Exon22 (Exon23 (1585b

FIG.4D

Appl. No. 09/720,086; 102(e): July 23, 2001

Dkt. No. 0609.4560002/JAG/KRM/DJN; Group Art Unit: 1642

Inventors: Li et al.; Tel: 202/371-2600

Title: De Novo DNA Cytosina Methyltransferase Genes, Polypeptides and Uses Thereof

	H	ΔI	IA
DNMT1 Dnmt1	DVFSGCGGLSEGFHQAG DVFSGCGGLSEGFHQAG	DVEMLCGGPPGQGFSGMNR DVEMLCGGPPCQGFSGMNR	YRPRFELLENVRNFVSFKR YRPRFELLENVRNFVSYRR
MET1(Ath)	DIFAGCGGLSHGLKKAG	QVDFINGGPPCQGFSGMNR	FRPRYFLLENVRTFVSFNK
Masc1	DTFCGGGGVSLGARQAG	HVDILHLSPPCQTFSRAHT	VRPRLFTVE ETDGIMDRQS
Masc.z Jumt.2	DIPAGCAGLICALDESA	CENDE I YOU'D TOUR DOUNK	YKRKFVLLENVKGLIIIKL KIDVVIIIENVKGEFVCCT
M.Spr	SLESGIGAFEAALRNIG	EFDLL VGGSPCQSFSVAGH	KQPKFFVFENVKGLINHDK
DNMT3A	SLFDGIATGLLVLKDLG	PFDLVIGGSPCNDLSIVNP	DRP FFWL FENVVAMGVSDK.
Dnmt3a	SLFDGIATGLLVLKDLG	66 S	DRPFFWLFENVVAMGVSDK
DNMT3B	SLFDGIATGYLVLKELG	PFDLVIGGSPCNDLSNVNP	DRPFFWMFENVVAMKVGDK
Dnmt3b	SLFDGIATGYLVLKELG	PFDLVIGGSPCNDLSNVNP	NRPFFWMFENVVAMKVNDK
Zmt3	SLFDGIATGYLVLRDLG	PFDLLIGGSPCNDLSIVNP	POPFFWLFENVTFMQTHVK
consensus	99	-N88N-	P-FEW
	XH	×	
DNMT1	RVVSVRECARSQGFP	LFGNILDKHRQVGNAVPPPLAKAIG	KAIG .
Dnmt1	RVVSVRECÄRSOGFP	FFGNILDRHRQVGNAVPPPLAKAIG	KAIG
MET1(Ath)	RILTVRECARSOGFP	FAGNINHKHROIGNAVPPPLAFALG	FALG
Masc1	RKFTVRELACIQGEP	FVGTLTDKRRIIGNAVPPPLSAAIM	AAIM
Masc2.	RVYTVRELARAQGEP	GLGGVKKWHRNIGNAVPVPLGEQIG	EQIG
Dnmt2	RYFTPKE I ANLOGEP	EKTTVKQRYR LL <mark>©</mark> NSLNVHVVAKL	AKLL
M.Spr	RRLTPLECFRLQAFD	AGISNSQLYKQTGNSITVTVLESIF	ESIF
DNMT3A	DILWCTEMERVFGFP	SNMSRLARQRLLGRSWSVPVIRHLF	RHLF
Dnmt3a	DILWCTEMERVFGFP	SNMSRLARORLLGRSWSVPVIRHLF	RHLF
DNMT3B	DVLWCTELERIFGFP	SNMGRGARQKLLGRSWSVPVIRHLF	RHLF
Dnmt3b	DVLWCTELERIFGFP	SNMGRGARQKLLGRSWSVPVIRHLF	
Zmt3	OHIWITELEKIFGEP	KSMGRPORGRVLGKSWSVPVIRHL	RHLL FIG. DA
consensus		dBR	

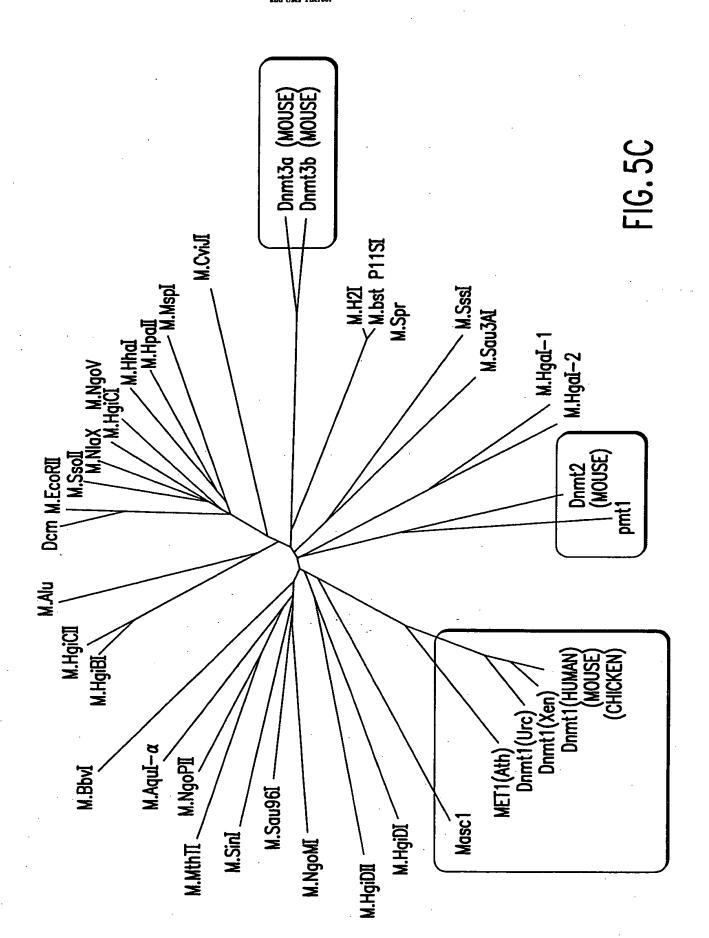
Appl. No. 09/720,086; 102(e): July 23, 2001
Dkt. No. 0609.4560002/JAG/KRM/DJN; Group Art Unit: 1642
Inventors: Li et al., Tel: 202/371-2600
Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

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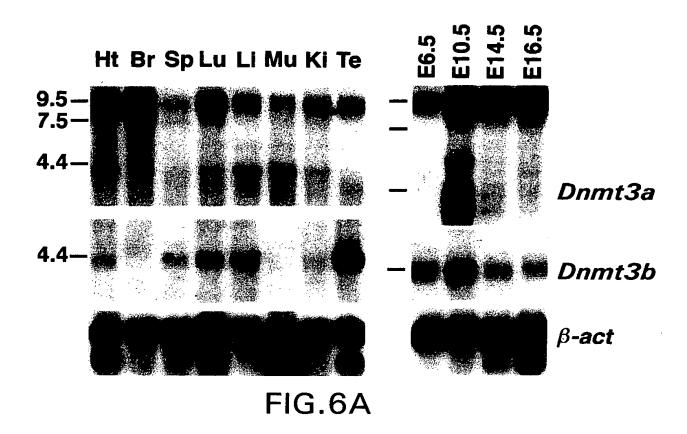
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DNMT3A Dnmt3a DNMT3B Dnmt3b Zmt3 ATRX Human ATRX Mouse Consensus	DNMT3A Dnmt3a DNMT3B Dnmt3b Zmt3 ATRX Human ATRX Mouse Consensus

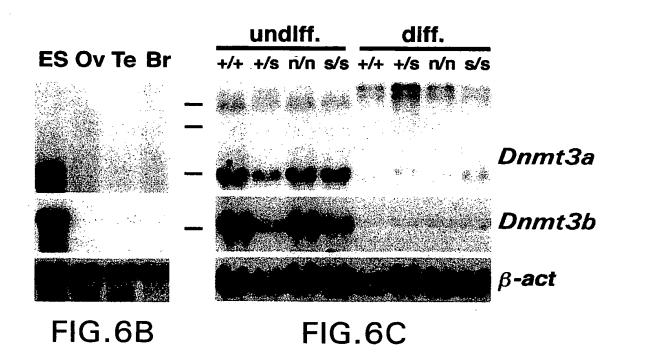
-IG.5B

Appl. No. 09/720,086, 102(e): July 23, 2001
Dkt. No. 0609 4560002/JAG/KRM/DJN; Group Art 1, 42
Inventors: Li et al.; Tel: 202/371-2600
Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

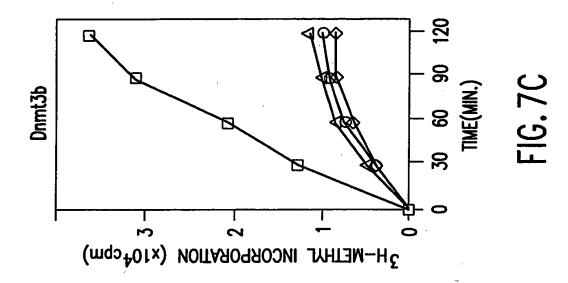


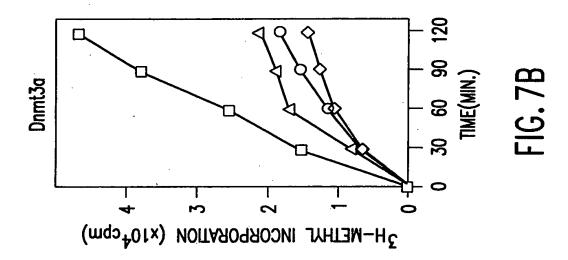
Appl. No. 09/720,086; 102(e): July 23, 2001
Dkt. No. 0609 4560002/JAG/KRM/DJN; Group Art Unit: 1642
[Inventors: Li et al.; Tel: 202/371-2600
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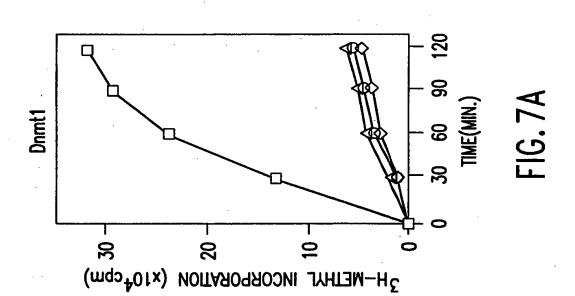




Appl. No. 09/720,086; 102(e): July 23, 2001
Dix. No. 0609.4560002/JAG/KRM/DJN; Group Art Unix: 164,
Inventors: Li et al.; Tel: 202/371-2600
Tüle: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides
and Uses There







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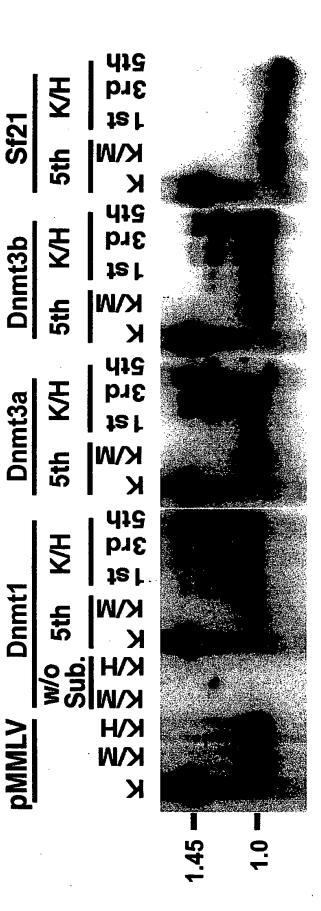
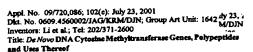
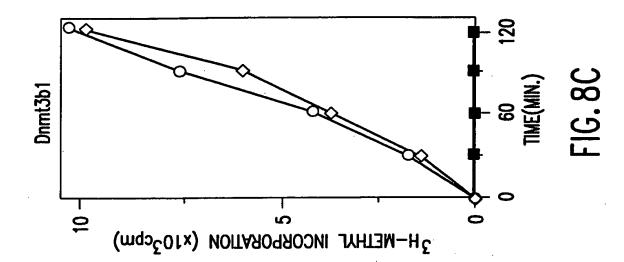
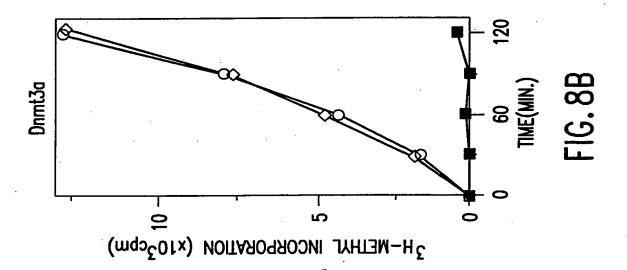
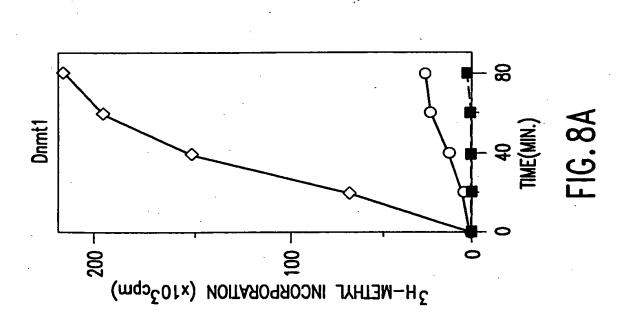


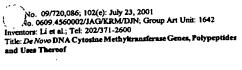
FIG.7D

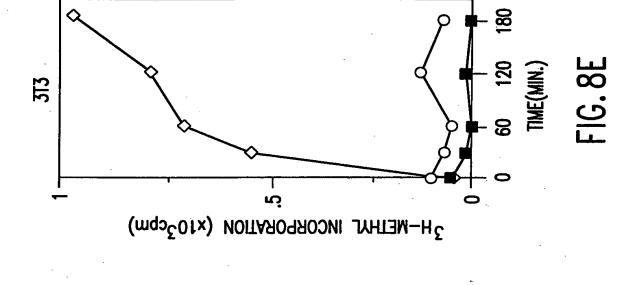








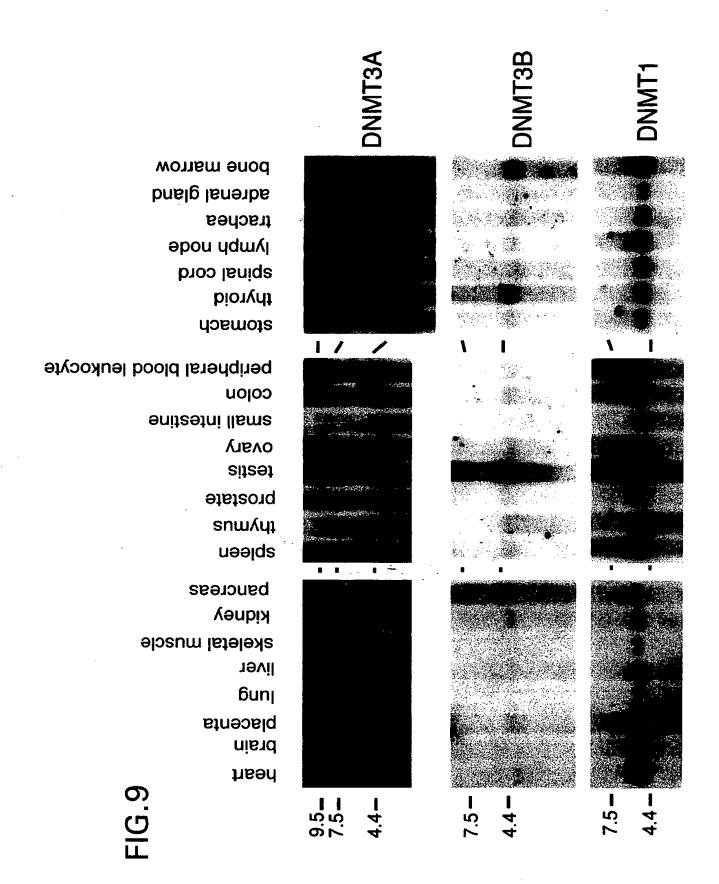




 3 H-METHYL INCORPORATION (x10 3 cpm)

Dnmt3b2

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Dict. No. 0609.4560002/JAG/KRM/DJN; Group Art Unit: 1642
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and Ulses Thereof



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chronic myelogenous leukemia K-562 colorectal adenocarcinoma SW480 lymphoblastic leukemia MOLT-4 promyelocytic leukemia HL-60 Burkiti's lymphoma Raji lung carcinoma A549 Hela cell S3

melanoma G361

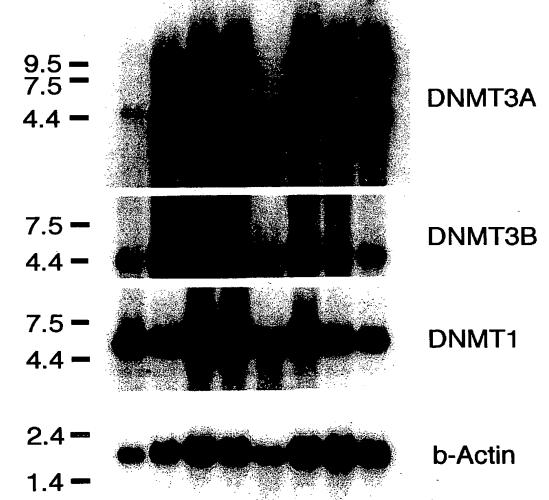


FIG.10